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                 patent numbers for U.S. applications
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                 patent records
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=> d 14 1-6 bib ab

- L4 ANSWER 1 OF 6 MEDLINE on STN
- AN 97312794 MEDLINE
- DN PubMed ID: 9169235
- TI Crystallization and preliminary X-ray analysis of neuropsin, a serine protease expressed in the limbic system of mouse brain.
- AU Kishi T; Kato M; Shimizu T; Kato K; Matsumoto K; Yoshida S; Shiosaka S; Hakoshima T
- CS Department of Molecular Biology, Nara Institute of Science and Technology (NAIST), Japan.
- SO Journal of structural biology, (1997 Apr) Vol. 118, No. 3, pp. 248-51. Journal code: 9011206. ISSN: 1047-8477.
- CY United States
- DT Journal; Article; (JOURNAL ARTICLE) (RESEARCH SUPPORT, NON-U.S. GOV'T)
- LA English
- FS Priority Journals
- EM 199706
- ED Entered STN: 9 Jul 1997
  Last Updated on STN: 3 Mar 2000
  Entered Medline: 20 Jun 1997
- AΒ Neuropsin (M(r) 25032) is a serine protease expressed in the limbic system of mouse brain. It has been implicated in various neurological processes including formation of memory and may be important as a drug target in the treatment of epilepsy. The recombinant protein was produced using a baculovirus expression system and was purified. Two crystal forms were obtained by a hanging-drop vapor-diffusion method with polyethylene glycol. Preliminary X-ray crystallographic analysis revealed that crystal form I belongs to triclinic space group P1 with unit cell dimensions a = 97.16 A, b = 97.12 A, c = 46.75 A and alpha = 99.17degrees, beta = 99.77 degrees, gamma = 117.35 degrees. Self-rotation function analysis of these data of form I indicates the position of a noncrystallographic threefold axis. There are six molecules in the crystallographic asymmetric unit. Crystal form II also belongs to triclinic space group P1 but has unit cell dimensions of a = 38.40 A, b =55.16 A, c = 65.37 A and alpha = 95.38 degrees, beta = 89.98 degrees, gamma = 110.46 degrees with two molecules in the crystallographic asymmetric unit. Form II has a noncrystallographic twofold axis. Intensity data to 3.1 A resolution for form I and to 2.2 A resolution for form II have been collected.
- L4 ANSWER 2 OF 6 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
- AN 2006:398347 BIOSIS
- DN PREV200600398661

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TI Methods and reagents for protease inhibition.
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- AU Albrecht, Hugo [Inventor]; Hengst, Ulrich [Inventor]; Monard, Denis [Inventor]
- CS Riehen, Switzerland
  ASSIGNEE: Novartis Forschungsstiftung Zweigniederlassung Friedrich
  Miescher Instittue for Biomedical Research
- PI US 07029877 20060418
- SO Official Gazette of the United States Patent and Trademark Office Patents, (APR 18 2006)
  CODEN: OGUPE7. ISSN: 0098-1133.
- DT Patent
- LA English
- ED Entered STN: 9 Aug 2006 Last Updated on STN: 9 Aug 2006
- AB There is provided a protease inhibitor and a method of inhibiting a protease selected from the group consisting of thrombin, chymotrypsin and neuropsin, by contacting the protease with an effective amount of a member of the phosphoethanolamine binding protein (PEBP) family.
- L4 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2006:1248277 CAPLUS
- DN 146:22551
- TI Random mutagenesis, screening and selection of protease variants with altered sensitivity to activity modulators
- IN Koltermann, Andre; Kettling, Ulrich; Haupts, Ulrich; Coco, Wayne; Tebbe, Jan; Votsmeier, Christian; Scheidig, Andreas
- PA Direvo Biotech AG, Germany
- SO Eur. Pat. Appl., 93pp. CODEN: EPXXDW
- DT Patent
- LA English

FAN.CNT 1

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PATENT NO.
                       KIND DATE
                                           APPLICATION NO.
                                                                  DATE
                                            _____
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                                          EP 2005-104543
    EP 1726643
PΙ
                         A1 20061129
                                                                    20050527
         R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
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             HR, LV, MK, YU
     US 20060269538
                                20061130
                                            US 2006-441635
                                                                    20060526
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     WO 2006125827
                         Α1
                                20061130
                                           WO 2006-EP62644
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR,
             KZ, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX,
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             VN, YU, ZA, ZM, ZW
         RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
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             GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
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     EP 1883696
                                20080206
                                           EP 2006-763303
                                                                    20060526
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             IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR
PRAI EP 2005-104543
                         Α
                                20050527
     US 2005-685566P
                          Ρ
                                20050527
     US 2005-686021P
                         Ρ
                                20050531
                          W
     WO 2006-EP62644
                                20060526
     The present invention provides a method for the selection of proteases
AB
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with altered sensitivity to one or more activity-modulating substances. The method combines the provision of a protease library (i.e., phage display library) encoding polynucleotide sequences generated by using PCR mutagenesis, expression of the enzymes, screening of the library in the presence of one or several activity-modulating substances, selection of variants with altered sensitivity to one or several activity-modulating substances and isolation of those polynucleotide sequences that encode for the selected variants. In particular, mutant variants of human trypsin are disclosed.

RE.CNT 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L4 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2006:684183 CAPLUS
- DN 146:2865
- ${
  m TI}$  Activation and enzymatic characterization of recombinant human kallikrein 8
- AU Kishi, Tadaaki; Cloutier, Sylvain M.; Kundig, Christoph; Deperthes, David; Diamandis, Eleftherios P.
- CS Department of Pathology and Laboratory Medicine, Mount Sinai Hospital, Toronto, ON, M5G 1X5, Can.
- SO Biological Chemistry (2006), 387(6), 723-731 CODEN: BICHF3; ISSN: 1431-6730
- PB Walter de Gruyter GmbH & Co. KG
- DT Journal
- LA English
- AΒ Human kallikrein 8 (hK8), whose gene was originally cloned as the human ortholog of a mouse brain protease, is known to be associated with diseases such as ovarian cancer and Alzheimer's disease. Recombinant human pro-kallikrein 8 was activated with lysyl endopeptidase-conjugated beads. Amino-terminal sequencing of the activated enzyme demonstrated the cleavage of a 9-aa propeptide from the pro-enzyme. The substrate specificity of activated hK8 was characterized using synthetic fluorescent substrates. HK8 showed trypsin-like specificity, as predicted from sequence anal. and enzymic characterization of the mouse ortholog. All synthetic substrates tested containing either arginine or lysine at P1 position were cleaved by hK8. The highest kcat/Km value of 20+103 M-1 s-1 was observed with Boc-Val-Pro-Arg-7-amido-4-methylcoumarin. The activity of hK8 was inhibited by antipain, chymostatin, and leupeptin. The concentration for 50% inhibition by the best inhibitor, antipain, was 0.46  $\mu M$ . The effect of different metal ions on the enzyme activity was analyzed. Whereas Na+ had no effect on hK8 activity, Ni2+ and Zn2+ decreased the activity and Ca2+, Mg2+, and K+ had a stimulatory effect. Ca2+ was the best activator, with an optimal concentration

of approx. 10  $\mu$ M.

RE.CNT 61 THERE ARE 61 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2005:1020554 CAPLUS
- DN 143:282218
- TI Protease activity assay method by using polymeric membrane
- IN Shiosaka, Sadao; Tamura, Hidenori
- PA Nara Institute of Science and Technology, Japan
- SO Jpn. Kokai Tokkyo Koho, 17 pp. CODEN: JKXXAF
- DT Patent
- LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE 		
PI PRAI	JP 2005253436 JP 2004-73625	A	20050922 20040315	JP 2004-73625	20040315		

AB A assay method for measuring protease (especially, neuropsin) activity with higher sensitivity and fewer sample amount than the traditional solution method. The method includes processes of (1) the sample containing protease is sticked to a polymeric membrane, (2) the protease is reacted with the substrate specific to the protease and (3) the signal resulted from the reaction is measured.

- L4 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2002:172125 CAPLUS
- DN 136:212778
- TI Identification of a novel brain serine protease inhibitory protein-phosphoethanolamine binding protein and methods and reagents for protease inhibition for the treatment of neurological disorders
- IN Albrecht, Hugo; Hengst, Ulrich; Monard, Denis
- PA Novartis Forschungsstiftung Zweigniederlassung Friedrich Miescher Institute for Biomedical Research, Switz.
- SO PCT Int. Appl., 39 pp. CODEN: PIXXD2
- DT Patent
- LA English

FAN CNT 1

F'AN.	PAT	PATENT NO.								APPLICATION NO.								
PI	WO	2002018623 2002018623			A2 20020307		0307	WO 2001-EP10043										
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		RW:	GH, DE,	GM, DK,	KE, ES,	LS, FI,	MW, FR,	MZ, GB, GA,	GR,	IE,	ΙΤ,	LU,	MC,	NL,	PT,	SE,	TR,	BF,
	AU	2420832 2002012184			A1 20020307 A 20020313			CA 2001-2420832 AU 2002-12184 EP 2001-980309						20010830 20010830				
			AT, IE,	BE, SI,	CH, LT,	DE, LV,	DK, FI,	ES, RO,	FR, MK,	GB, CY,	GR, AL,	IT, TR	LI,	LU,	NL,	SE,		PT,
	US US	3 7029877 3 20060177432			B2 20060418 A1 20060810			US 2003-362642 US 2005-311974										
PKAI	WO	2000-21497 2001-EP10043 2003-362642			W		20010830											

AB The present invention is based on the discovery of a novel serine protease inhibitory protein-phosphoethanolamine binding protein (PEBP). PEPB is identified by the detection of a novel thrombin inhibitory activity in the brain of protease nexin-1(-/-) mice, a gene knockout for the only known endogenous protease inhibitor protease nexin-1 that specifically interferes with thrombotic activity and is expressed in the brain. PEBP exerts inhibitory activity against several serine proteases including thrombin, neuropsin, and chymotrypsin, whereas trypsin, tissue type plasminogen activator, and elastase are not affected. PEBP immunoreactivity is found on the surface of Rat-1 fibroblast cells and although its sequence contains no secretion signal, PEBP-H6 can be

purified from the conditioned medium upon recombinant expression. The method of inhibiting a protease selected from the group consisting of thrombin, chymotrypsin and neuropsin, by contacting the protease with an effective amount of PEBP.